

### Amendments to the Specification

Please replace the paragraph at page 10, lines 7-15 with the following paragraph:

**FIGURE 14.** Potential functional domains of TARP. (A) TARP contains a potential leucine zipper motif and phosphorylation sites. A potential leucine zipper motif is indicated with boxed leucines followed by a basic region that is underlined. cAMP- and cGMP-dependent protein kinase phosphorylation sites (amino acids 46-49 and 55-58, see SEQ ID NO: 14) and protein kinase C phosphorylation sites (amino acids 19-21 and 20-22, see SEQ ID NO: 15) are outlined. (B) Protein sequence comparison of TARP with Tup1. Amino acid sequences for TARP (42-57, SEQ ID NO: 16), *Dictyostelium discoideum* Tup1 (dTup1, 521-536, SEQ ID NO: 17) and *Saccharomyces cerevisiae* Tup1 (yTup1, 626-660, SEQ ID NO: 18) are shown. Conserved residues are boxed.

Please replace the paragraph at page 13, lines 9-28 with the following paragraph:

To determine if TARP shares homology with any known proteins, we performed a protein BLAST search against ~~GenBank~~ GENBANK® (database). This search indicated that the amino acid sequence of TARP shares some homology to *Dictyostelium discoideum* Tup1 (~~GenBank~~ GENBANK® (database) accession no. AAC29438) and *Saccharomyces cerevisiae* Tup1 (Williams, F. E. *et al.*, *Mol. Cell. Biol.* 10:6500-6511 (1990)) (Figure 14B). Yeast Tup1 is normally found in a complex with Cyc8(Ssn6) and is required for transcriptional repression of genes that are regulated by glucose, oxygen and DNA damage (Tzamarias, D. *et al.*, *Genes Dev.* 9:821-831 (1995)). Neither Cyc8(Ssn6) nor Tup1 binds DNA, but each acts as a part of a corepressor complex through interactions with specific DNA-binding proteins such as  $\alpha 2$ , Mig1, Rox1 and a1 (Tzamarias, D. *et al.*, *Genes Dev.* 9:821-831 (1995)). The C'-terminal half of Tup1 contains six repeats of a 43-amino acid sequence rich in aspartate and tryptophan, known as WD-40 or  $\beta$ -transducin repeats (Williams, F. E. *et al.*, *Mol. Cell. Biol.* 10:6500-6511 (1990); Fong, H. K. *et al.*, *Proc. Natl. Acad. Sci. USA* 83:2162-2166 (1986)). WD-40 repeats have been identified in many proteins and play a role in protein-protein interactions. Importantly, Tup1 has been shown to interact with  $\alpha 2$  through two of its WD-40 repeats (Komachi, K. *et al.*, *Genes Dev.*

8:2857-2867 (1994)). It is interesting to note that TARP shares homology with the fifth WD-40 repeat of Tup1 (Figure 14B 7C). Because TARP is a nuclear protein, its homology with Tup1 suggests that TARP is a member of a functional nuclear protein complex involved in transcriptional regulation.

Please replace the paragraph at page 42, line 31 to page 43, lines 18 with the following paragraph:

RNA dot blot (RNA master blot, Clontech, Palo Alto, CA), and Northern blot (MTN, Clontech, Palo Alto, CA), were performed on a variety of human tissues. Northern blot was also performed on mRNA from prostate adenocarcinoma cell lines, LNCaP and PC-3 (ATCC, Rockville, MD). Isolation of poly(A) RNA was carried out using the FASTTRACK™ ~~FastTrack~~ (kit) (Invitrogen, Carlsbad, CA). RNA was electrophoresed on a 1% agarose gel and transferred to nylon-based membranes (GENESCREEN PLUS™ ~~GeneScreen Plus~~, DuPont, Wilmington, DE), according to established procedures. Ausubel, *supra*. A cDNA probe specific for the untranslated 3' end (3' UTR) of the TCR $\gamma$  transcript was made from EST plasmid ng79d11 (Genome Systems, St. Louis, MO). A probe specific for the constant domain of the TCR $\gamma$  transcript (TCR C $\gamma$ ) was made from LNCaP cDNA and a probe for the constant domain of the TCR $\delta$  transcript (TCR C $\delta$ ) was made from a TCR $\delta$  plasmid. A human  $\beta$ -actin probe was used as a quantity control of the mRNA preparations. Probes were labeled with  $^{32}\text{P}$  by random primer extension (Lofstrand Labs Limited, Gaithersburg, MD) to a specific activity of 1  $\mu\text{Ci}/\text{ng}$ . The RNA membranes were blocked for 2 hours at 45° C in hybridization solution containing 50% formamide (Hybrisol I, Oncor, Gaithersburg, MD) and then probed for 15 hours at 45° C with 20  $\mu\text{Ci}$  cDNA in 20 ml hybridization solution. The membranes were washed twice for 15 minutes at room temperature in 2xSSC/0.1%SDS and twice for 20 minutes at 55-65° C in 0.1%SSC/0.1%SDS. The membranes were exposed to an imaging film (X-OMAT™, Kodak, Rochester, NY) at -80° C before development.

Please replace the paragraph at page 47, lines 3-11 with the following paragraph:

TCR  $\gamma$ -chain protein is normally co-expressed with the TCR  $\delta$ -chain protein. Since the TCR $\gamma$  gene is transcriptionally active in human prostate, we went on to analyze the transcriptional activity of the TCR $\gamma$  gene. The dbEST was analyzed (<http://www.ncbi.nlm.nih.gov/BLAST> on the NCBI website) using the TCR $\delta$  transcript nucleotide sequence. ESTs from prostate cDNA libraries did not match any part of the TCR  $\delta$ -chain transcript. Furthermore, Northern blot analysis did not detect any prostate expression of TCR $\delta$  mRNA, Figure 3B (lane 3). We conclude that the TCR $\delta$  gene is silent in prostate. As expected, TCR $\delta$  transcripts are expressed in spleen, thymus and blood leukocytes, Figure 3B.

Please replace the paragraph found at page 59, line 25-35 with the following paragraph:

The TARP antibody recognizes a doublet in prostate and breast nuclear extracts (Figure 13A). The faster 7 kDa band comigrates with the His-TARP recombinant protein, while the weaker band runs at a larger molecular weight. One possible explanation for the 9 kDa band is post-translational modifications. To determine if TARP contains any known post-translational modification sites, we analyze the TARP amino acid sequence using the PROSITE program of the *Swiss Institute of Bioinformatics* ExPASy proteomics server [<http://www.expasy.ch>] available on the internet] (Appel, R.D. *et al.*, *Trends Biochem. Sci.* **19**:248-260 (1994); Hofmann, K *et al.*, *Nucleic Acids Res.* **27**:215-219 (1999)). As shown in Figure 14A, many potential phosphorylation sites were found including cAMP- and cGMP-dependent protein kinase phosphorylation sites (RRAT (SEQ ID NO:32) and RRGTT (SEQ ID NO:33)) and

Please replace the paragraph found at page 59, lines 4-24 with the following paragraph:

To determine if TARP shares homology with any known proteins, we performed a protein BLAST search against ~~GenBank~~ GENBANK® (database). This search indicated that the amino acid sequence of TARP shares some homology to *Dictyostelium discoideum* Tup1 (~~GenBank~~ GENBANK® (database) accession no. AAC29438) and *Saccharomyces cerevisiae* Tup1 (Williams, F. E. *et al.*, *Mol. Cell. Biol.* **10**:6500-6511 (1990)) (Figure 7C-14B). Yeast

Tup1 is normally found in a complex with Cyc8(Ssn6) and is required for transcriptional repression of genes that are regulated by glucose, oxygen and DNA damage (Tzamarias, D. *et al.*, *Genes Dev.* 9:821-831 (1995)). Neither Cyc8(Ssn6) nor Tup1 binds DNA, but each acts as a part of a corepressor complex through interactions with specific DNA-binding proteins such as  $\alpha 2$ , Mig1, Rox1 and a1 (Tzamarias, D. *et al.*, *Genes Dev.* 9:821-831 (1995)). The C'-terminal half of Tup1 contains six repeats of a 43-amino acid sequence rich in aspartate and tryptophan, known as WD-40 or  $\beta$ -transducin repeats (Williams, F. E. *et al.*, *Mol. Cell. Biol.* 10:6500-6511 (1990); Fong, H. K. *et al.*, *Proc. Natl. Acad. Sci. USA* 83:2162-2166 (1986)). WD-40 repeats have been identified in many proteins and play a role in protein-protein interactions. Importantly, Tup1 has been shown to interact with  $\alpha 2$  through two of its WD-40 repeats (Komachi, K. *et al.*, *Genes Dev.* 8:2857-2867 (1994)). It is interesting to note that TARP shares homology with the fifth WD-40 repeat of Tup1 (Figure 14B 7C). Because TARP is a nuclear protein, its homology with Tup1 suggests that TARP may be a member of a functional nuclear protein complex involved in transcriptional regulation. Therefore, it is necessary to identify TARP-interacting proteins in order to determine its function.